(FILE 'HOME' ENTERED AT 13:34:05 ON 30 OCT 2002)

	FILE 'BIOSIS, CABA, CAPLUS, EMBASE, LIFESCI, MEDLINE, SCISEARC	Η,
	USPATFULL, JAPIO' ENTERED AT 13:34:24 ON 30 OCT 2002	
L1	12459 S FOOT AND MOUTH DISEASE VIRUS	
L2	3177 S L1 AND VACCINE	
L3	16405 S L2 AND PILI OR FIMBRIAE	
L4	1018 S L3 AND (FUSION OR HYBRID OR CHIMERIC)	
L5	0 S L4 AND FOREGIN EPITOPES	
L6	44 S L4 AND FOREIGN EPITOPES	
T.7	21 DUP REM I.6 (23 DUPLICATES REMOVED)	

7 ANSWER 1 OF 21 USPATFULL

AB In order to obtain a novel binding protein against a chosen target, DNA molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein.

AN 2002:272761 USPATFULL

TI Directed evolution of novel binding proteins

IN Ladner, Robert Charles, Ijamsville, MD, UNITED STATES
Guterman, Sonia Kosow, Belmont, MA, UNITED STATES
Roberts, Bruce Lindsay, Milford, MA, UNITED STATES
Markland, William, Milford, MA, UNITED STATES
Ley, Arthur Charles, Newton, MA, UNITED STATES
Kent, Rachel Baribault, Boxborough, MA, UNITED STATES

PI US 2002150881 A1 20021017 AI US 2001-781988 A1 20010214 (9)

RLI Continuation of Ser. No. US 1998-192067, filed on 16 Nov 1998, ABANDONED Continuation of Ser. No. US 1995-415922, filed on 3 Apr 1995, PATENTED Continuation of Ser. No. US 1993-9319, filed on 26 Jan 1993, PATENTED Division of Ser. No. US 1991-664989, filed on 1 Mar 1991, PATENTED Continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, ABANDONED Continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, ABANDONED

PRAI WO 1989-US3731 19890901

DT Utility

FS APPLICATION

LREP BROWDY AND NEIMARK, P.L.L.C., 624 Ninth Street, N.W., Washington, DC, 20001

CLMN Number of Claims: 18 ECL Exemplary Claim: 1 DRWN 16 Drawing Page(s)

LN.CNT 15696

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 2 OF 21 USPATFULL

AB A strategically modified hepatitis B core protein is described, where an insert is provided, preferably in an immunodominant region of the nucleocapsid protein, containing a chemically reactive amino acid residue. The modified hepatitis B core protein or its aggregated nucleocapsid protein particles can be pendently linked to a hapten to form a modified nucleocapsid conjugate. Such a conjugate is useful in the preparation of vaccines or antibodies. The modified hepatitis B core protein can also be modified to include a T cell epitope.

AN 2001:71101 USPATFULL

TI Strategically modified hepatitis B core proteins and their derivatives

IN Birkett, Ashley J., Solana Beach, CA, United States

PA Immune Complex Corporation, San Diego, CA, United States (U.S. corporation)

PI US 6231864 B1 20010515 AI US 1999-248588 19990211 (9) PRAI US 1998-74537P 19980212 (60) DT Utility FS Granted

EXNAM Primary Examiner: Wortman, Donna C.

LREP Welsh & Katz, Ltd.
CLMN Number of Claims: 22
ECL Exemplary Claim: 1

DRWN 1 Drawing Figure(s); 1 Drawing Page(s)

LN.CNT 1665

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- L7 ANSWER 3 OF 21 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE
 1
- Based on the prediction of the hydrophilicity, epitopes, secondary structure and flexibility of the CS3 subunit, a novel vector pCSX72 which permits the insertion of foreign epitopes into CS3 at the position of 72nd aa was constructed. Two epitopes, the VP1 of FMDV and a ten-peptides epitope of C-myc, were displayed with it respectively. Compared with the two previously-constructed vectors, the vector pCSX72 expressed the hybrid fimbriae in higher level. Mice produced dual immune response against the CS3 and the inserted epitopes when they were immunized by injecting the live recombinant bacteria intraperitoneally.
 - AN 2002:99199 BIOSIS
 - DN PREV200200099199
 - TI Construction of a novel display vector deriving from CS3 **fimbriae** of human enterotoxigenic Escherichia coli.
 - AU Gao Rong-Kai; Zhang Zhao-Shan (1); Li Shu-Qin; Huang Cui-Fen
 - CS (1) Institute of Biotechnology, Academy of Military Medical Sciences, Beijing, 100071: Zhangzs@nic.bmi.ac.cn China
 - SO Acta Genetica Sinica, (Oct., 2001) Vol. 28, No. 10, pp. 971-980. print. ISSN: 0379-4172.
 - DT Article
 - LA English
 - L7 ANSWER 4 OF 21 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 2
 - AB The gene fragments coding for C3 epitope of poliovirus and a ten-peptides epitope of C-myc were synthesized and inserted into pCSB136 and pCSX72 resp. to confirm the possibility of pCSB136 and pCSX72 as vectors for displaying heterologous epitopes. The recombinants were screened by whole-strain PCR. The expression of recombinant proteins were detected by whole-cell ELISA and electronic microscopy. The results indicated the recombinant proteins were expressed as hybrid fimbriae, and the antigenicity of both CS3 and inserted epitopes kept. All results above showed vectors pCSB136 and pCSX72 could be used to display the foreign epitopes.
 - AN 2001:825914 CAPLUS
 - DN 137:77480
 - TI Expression of C3 epitope of poliovirus and a ten-peptides epitope of C-myc on surface of recombinant bacteria
 - AU Gao, Rongkai; Zhang, Zhaoshan; Li, Shuqin; Huang, Cuifen
 - CS Beijing Institute of Biotechnology, Beijing, 100071, Peop. Rep. China
 - SO Shengwu Gongcheng Xuebao (2001), 17(5), 539-542 CODEN: SGXUED; ISSN: 1000-3061
 - PB Kexue Chubanshe
 - DT Journal
 - LA Chinese
 - L7 ANSWER 5 OF 21 CAPLUS COPYRIGHT 2002 ACS
 - AB A method of generating **chimeric** genes encoding a **fusion** product of the agfA fimbrin and a foreign protein, such as an antigen, in a Salmonella host by chromosomal gene replacement is described. One embodiment of the invention is exemplified by the expression of a model epitope (PT3) obtained from the GP63 protein of Leishmania major, by formation of recombinant agfA genes encoding PT3 fusing proteins

recombined at 10 different sites throughout the agfA gene. fusions are shown to be expressed in the thin aggregative fimbriae on the surface of bacterial cell. The AgfA fimbrin of Salmonella (CsgA for E. coli) provides a flexible and stable vehicle for the expression of foreign epitopes in enterobacteriaceae and the subsequent thin aggregative fimbrae (curli) expression product provide an ideal organelle for presentation of the foreign epitopes at the cell surface. AN 2000:725786 CAPLUS DN 133:306338 Use of the agfA fimbrin of Salmonella to present foreign proteins on the TIsurface of a bacterial host IN White, Aaron P.; Doran, James L.; Collison, S. Karen; Kay, William W. PA Innovation and Development Corporation, University of Victoria, Can. so PCT Int. Appl., 139 pp. CODEN: PIXXD2 DT Patent LA English FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE ----_____ -----WO 2000060102 WO 2000-CA356 20000405 PΙ A2 20001012 WO 2000060102 **A**3 20010104 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG PRAI US 1999-127888P 19990405 Р ANSWER 6 OF 21 USPATFULL 1.7 A CS31A protein capsule subunit having an aminoacid sequence modified by AB at least one heterologous peptide, the CS31A protein capsule comprising said subunit, and micro-organisms having the CS31A protein capsule with its subunit aminoacid sequence modified by at least one heterologous peptide, are disclosed. Methods for preparing said subunits, CS31A protein capsules comprising same, and micro-organisms having CS31A protein capsules, as well as the use thereof for preparing vaccines, producing peptides and preparing immunoassays, are also disclosed. 2000:98007 USPATFULL AN ΤI ClpG subunit of CS31A protein capsule containing heterologous peptides Girardeau, Jean-Pierre, Saint Genes Champanelle, France IN Martin, Christine, La Roche Blanche, France Mechin, Marie-Claire, Beaumont, France Der Vartanian, Maurice, Saint Genes Champanelle, France Bousquet, Fran.cedilla.ois, Ceyrat, France PA Institut National de la Recherche Agronomique-INRA, Paris, France (non-U.S. corporation) US 6096321 PΙ 20000801 WO 9414967 19940707 US 1996-491954 AΙ 19960216 (8) WO 1993-FR1281 19931221 19960216 PCT 371 date 19960216 PCT 102(e) date PRAI FR 1992-15464 19921222 DTUtility Granted FS EXNAM Primary Examiner: Chin, Christopher L.; Assistant Examiner: Ryan, V. LREP Schnader Harrison Segal & Lewis LLP CLMN Number of Claims: 29

ECL Exemplary Claim: 1 61 Drawing Figure(s); 53 Drawing Page(s) CAS INDEXING IS AVAILABLE FOR THIS PATENT. L7 ANSWER 7 OF 21 USPATFULL The present invention is concerned with vaccination of mammals against AB GnRH. The vaccine comprises a GnRH peptide conjugate to E. coli fimbrial-filaments and elicits an immune response against GnRH. 2000:12446 USPATFULL AN Carrier system against GnRH ΤI Van Der Zee, Anna, Woerden, Netherlands IN Van Die, Irma Marianne, Gouda, Netherlands Hoekstra, Willem Pieter Martin, Zeist, Netherlands Gielen, Josephus Theodorus, St. Antohonis, Netherlands PA Akzo Nobel N.V., Arnhem, Netherlands (non-U.S. corporation) PΙ US 6019983 20000201 ΑI US 1995-521079 19950829 (8) Continuation of Ser. No. US 1993-78661, filed on 16 Jun 1993, now RLI abandoned PRAI NL 1982-92201775 19820619 DT Utility FS Granted Primary Examiner: Sidberry, Hazel F. EXNAM Gormley, Mary E., Blackstone, William M. LREP CLMN Number of Claims: 6 Exemplary Claim: 1 ECL DRWN 9 Drawing Figure(s); 9 Drawing Page(s) LN CNT 1366 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- L7 ANSWER 8 OF 21 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE
- AB Recombinant live oral vaccines expressing pathogen-derived antigens offer a unique set of attractive properties. Among these are the simplicity of administration, the capacity to induce mucosal and systemic immunity, and the advantage of permitting genetic manipulation for optimal antigen presentation. In this study, the benefit of having a heterologous antigen expressed on the surface of a live vector rather than intracellularly was evaluated. Accordingly, the immune response of mice immunized with a Salmonella enterica serovar Typhimurium vaccine strain expressing the Escherichia coli 987P fimbrial antigen on its surface (Fas+) was compared with the expression in the periplasmic compartments (Fas-). Orally immunized BALB/c mice showed that 987P fimbriated Salmonella serovar Typhimurium CS3263 (aroA asd) with pCS151 (fas+ asd+) elicited a significantly higher level of 987P-specific systemic immunoglobulin G (IgG) and mucosal IgA than serovar Typhimurium CS3263 with pCS152 (fasD mutant, asd+) expressing 987P periplasmic antigen. Further studies were aimed at determining whether the 987P fimbriae expressed by serovar Typhimurium chi4550 (cya crp asd) could be used as carriers of foreign epitopes. For this, the vaccine strain was genetically engineered to express chimeric fimbriae carrying the transmissible qastroenteritis virus (TGEV) C (379-388) and A (521-531) epitopes of the spike protein inserted into the 987P major fimbrial subunit FasA. BALB/c mice administered orally serovar Typhimurium chi4550 expressing the chimeric fimbriae from the tet promoter in pCS154 (fas+ asd+) produced systemic antibodies against both fimbria and the TGEV C epitope but not against the TGEV A epitope. To improve the immunogenicity of the chimeric fimbriae, the in vivo inducible nirB promoter was inserted into pCS154, upstream of the fas genes, to create pCS155. In comparison with the previously used vaccine, BALB/c mice immunized orally with serovar Typhimurium chi4550/pCS155 demonstrated significantly higher levels of serum IqG and mucosal IgA against 987P fimbria. Moreover, mucosal IgA against the TGEV C

epitope was only detected with serovar Typhimurium chi4550/pCS155. The induced antibodies also recognized the epitopes in the context of the full-length TGEV spike protein. Hence, immune responses to heterologous chimeric fimbriae on Salmonella vaccine vectors can be optimized by using promoters known to be activated in vivo.

AN 2000:291353 BIOSIS

DN PREV200000291353

- TI Mucosal and systemic immune responses to **chimeric fimbriae** expressed by Salmonella enterica serovar Typhimurium vaccine strains.
- AU Chen, Huaiqing; Schifferli, Dieter M. (1)
- CS (1) University of Pennsylvania School of Veterinary Medicine, 3800 Spruce St., Philadelphia, PA, 19104-6049 USA
- SO Infection and Immunity, (June, 2000) Vol. 68, No. 6, pp. 3129-3139. print. ISSN: 0019-9567.
- DT Article
- LA English
- SL English
- L7 ANSWER 9 OF 21 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 4
- AB Objective: To construct the display vector based on the CS3 pili of enterotoxigenic Escherichia coli. Methods: The secondary structure antigen epitopes, hydrophilicity and flexibility of CS3 subunit were predicted with the Goldkey software. Based on the prediction, the site for inserting heterologous epitopes was chosen. Mutation was done using the overlapping extention PCR. The gene fragment coding for the VP1 of foot-mouth disease virus (FMDV) was synthesized and inserted into CS3. The surface expression of hybrid protein was examined using whole-cell ELISA, electron microscopy and immuno-electron microscopy. Mice were immunized by injecting the recombinant bacteria intraperitoneally to evaluate the immunogenicity of the hybrid proteins. Results: The VP1 of FMDV was displayed on the surface of the recombinant cells. The fusion proteins were expressed as hybrid pili. Mice produced antibody response against CS3 and the VP1 of FMDV. Conclusion: The CS3 pili can be a vector to express the foreign epitopes on the surface of the recombinant cells, and it may probably be an expression vector for the construction of the live gene engineering vaccine.
- AN 2001:49887 BIOSIS
- DN PREV200100049887
- TI Construction of a display vector based on the CS3 pili of enterotoxigenic Escherichia coli.
- AU Gao Rongkai; Zhang Zhaoshan (1); Li Shuqin
- CS (1) Academy of Military Medical Science, Institute of Biotechnology, Beijing, 100071: zhangzs@nic.bmi.ac.cn China
- SO Zhonghua Weishengwuxue He Mianyixue Zazhi, (November, 2000) Vol. 20, No. 6, pp. 485-488. print. ISSN: 0254-5101.
- DT Article
- LA Chinese
- SL Chinese; English
- L7 ANSWER 10 OF 21 SCISEARCH COPYRIGHT 2002 ISI (R)
- The display of peptide segments on the surface of bacteria offers many new and exciting applications in biotechnology and medical research. Fimbria-assisted display of heterologous sequences is a paradigm for chimeric organelle display on bacteria. Fimbriac are particularly attractive candidates for epitope display for several reasons: (1) they are present in extremely high numbers at the cell surface, (2) they are strong immunogens, (3) they possess inherent adhesive properties, and (4) they can be easily purified. The majority of work dealing with fimbria-assisted peptide display has been focused on the development of

recombinant vaccines. A number of different fimbrial types have been used to display immune-relevant sectors of various foreign proteins. Chimeric fimbrial vaccines can be used in the context of purified proteins, however the potential also exists to exploit this technology for the development of live recombinant vaccines. Work has also bern performed demonstrating the amenability of fimbriae towards the powerful technology of random peptide display. This review summarises the current state of research in this field.

AN 2000:632511 SCISEARCH

GA The Genuine Article (R) Number: 344MZ

TI **Fimbriae**-assisted bacterial surface display of heterologous peptides

AU Klemm P (Reprint); Schembri M A

CS TECH UNIV DENMARK, DEPT MICROBIOL, BLDG 301, DK-2800 LYNGBY, DENMARK (Reprint)

CYA DENMARK

SO INTERNATIONAL JOURNAL OF MEDICAL MICROBIOLOGY, (JUL 2000) Vol. 290, No. 3, pp. 215-221.

Publisher: UBBAN & FISCHER VERLAG BRANCH OFFICE JENA D.O. BOY 100527

Publisher: URBAN & FISCHER VERLAG, BRANCH OFFICE JENA, P O BOX 100537, D-07705 JENA, GERMANY.

ISSN: 1438-4221.

DT Article; Journal

FS LIFE

LA English

REC Reference Count: 41
ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

- L7 ANSWER 11 OF 21 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 5
- AB The strong immunogenicity of bacterial fimbriae results from their polymeric and proteinaceous nature, and the protective role of these immunogens in experimental or commercial vaccines is associated with their capacity to induce antiadhesive antibodies. Fimbria-mediated intestinal colonization by enteropathogens typically leads to similar antibody responses. The possibility of taking advantage of these properties was investigated by determining whether enteroadhesive fimbriae, like the 987P fimbriae of enterotoxigenic Escherichia coli, can serve as carriers for foreign antigens without losing their adhesive characteristics. Random linker insertion mutagenesis of the fasA gene encoding the major 987P subunit identified five different mutants expressing wild-type levels of fimbriation. The linker insertion sites of these mutants were used to introduce three continuous segments of viral surface glycoproteins known to be accessible to antibodies. These segments encode residues 11 to 19 or 272 to 279 of herpes simplex virus type 1 (HSV-1) glycoprotein D (gD(11-19) and gD(272-279), respectively) or residues 379 to 388 of the transmissible gastroenteritis virus (TGEV) spike protein (S(379-388)). Studies of bacteria expressing fimbriae incorporating mutated FasA subunits alone or together with wild-type FasA subunits (hybrid fimbriae) indicated that foreign epitopes were best exported and displayed on assembled fimbriae when they were inserted near the amino terminus of FasA. Fimbriated bacteria expressing FasA subunits carrying the HSV gD(11-19) or the TGEV S(379-388) epitope inserted between the second and third residues of mature FasA elicited high levels of foreign epitope antibodies in all rabbits immunized parenterally. Antibodies against the HSV epitope were also shown to recognize the epitope in the context of the whole gD protein. Because the 987P adhesive subunit FasG was shown to be present on mutated fimbriae and to mediate bacterial attachment to porcine intestinal receptors, polymeric display of foreign epitopes on 987P offers new opportunities to test the potential beneficial effect of enteroadhesion for mucosal immunization and protection against various enteric pathogens.

AN 1999:99340 BIOSIS

DN PREV199900099340

Polymeric display of immunogenic epitopes from herpes simplex virus and TI transmissible gastroenteritis virus surface proteins on an enteroadherent Rajini Rani, D. B.; Bayer, Manfred E.; Schifferli, Dieter M. (1) ΑU (1) Univ. Pa. Sch. Veterinary Med., 3800 Spruce St., Philadelphia, PA CS 19104-6049 USA Clinical and Diagnostic Laboratory Immunology, (Jan., 1999) Vol. 6, No. 1, SO pp. 30-40. ISSN: 1071-412X. DT Article English LA L7 ANSWER 12 OF 21 USPATFULL AB In order to obtain a novel binding protein against a chosen target, DNA molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein. AN 1998:143904 USPATFULL ΤI Directed evolution of novel binding proteins IN Ladner, Robert Charles, Ijamsville, MD, United States Gutterman, Sonia Kosow, Belmont, MA, United States Roberts, Bruce Lindsay, Milford, MA, United States Markland, William, Milford, MA, United States Ley, Arthur Charles, Newton, MA, United States Kent, Rachel Baribault, Boxborough, MA, United States PA Dyax, Corp., Cambridge, MA, United States (U.S. corporation) PΙ US 5837500 19981117 ΑI US 1995-415922 19950403 (8) Continuation of Ser. No. US 1993-9319, filed on 26 Jan 1993, now RLI patented, Pat. No. US 5403484 which is a division of Ser. No. US 1991-664989, filed on 1 Mar 1991, now patented, Pat. No. US 5223409 which is a continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned which is a continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, now abandoned DΤ Utility FS Granted Primary Examiner: Ulm, John Cooper, Iver P. EXNAM LREP CLMN Number of Claims: 43 ECL Exemplary Claim: 1 DRWN 16 Drawing Figure(s); 16 Drawing Page(s) LN.CNT 15973 CAS INDEXING IS AVAILABLE FOR THIS PATENT. L7 ANSWER 13 OF 21 USPATFULL AB The present invention is concerned with vaccination of mammals against GnRH. The vaccine comprises a GnRH peptide conjugate to E. coli fimbrial-filaments and elicits an immune response against GnRH.

ΑN

TТ

IN

97:101896 USPATFULL

Carrier system against GNRH

Van Der Zee, Anna, Woerden, Netherlands

Van Die, Irma Marianne, Gouda, Netherlands Hoekstra, Willem Pieter Martin, Zeist, Netherlands Gielen, Josephus Theodorus, St. Antohonis, Netherlands AKZO Nobel N.V., Arnhem, Netherlands (non-U.S. corporation) PA PΙ US 5684145 19971104 ΑI US 1995-453588 19950530 (8) Division of Ser. No. US 1993-78661, filed on 16 Jun 1993, now abandoned RLI NL 1992-1775 PRAI 19920618 DT Utility Granted FS EXNAM Primary Examiner: Sidberry, Hazel F. Gormley, Mary E. Number of Claims: 8 ECL Exemplary Claim: 1 9 Drawing Figure(s); 9 Drawing Page(s) LN.CNT 1299 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 14 OF 21 SCISEARCH COPYRIGHT 2002 ISI (R) 1.7 The potential of the major structural protein of type 1 AB fimbriae as a display system for heterologous sequences was tested. As a reporter-epitope, a heterologous sequence mimicking a neutralizing epitope of the cholera toxin B chain was inserted, in one or two copies, into four different positions in the fimA gene. This was carried out by introduction of new restriction sites by PCR-mediated site-directed mutagenesis of fimA in positions predicted to correspond to optimally surface-located regions of the subunit protein. Subsequently, the synthetic cholera-toxin-encoding DNA segment was inserted. Several of the chosen positions seemed amenable even for large foreign inserts; the chimeric proteins were exposed on the bacterial surface and the cholera toxin epitope was authentically displayed, i.e. it was recognized on bacteria by specific antiserum. Display of chimeric fimbriae was tested with respect to host background in three different Escherichia coli strains, i.e. an isogenic set of K-12 strains, differing in the presence of an indigenous fim gene cluster, as well as a wild-type isolate. Immunization of rabbits with purified chimeric fimbriae resulted in serum which specifically recognized cholera toxin B chain, confirming the utility of the employed strategy. ΑN 97:462137 SCISEARCH GA The Genuine Article (R) Number: XE251 TI Authentic display of a cholera toxin epitope by chimeric type 1 fimbriae: Effects of insert position and host background ΑU StentebjergOlesen B; Pallesen L; Jensen L B; Christiansen G; Klemm P (Reprint) CS TECH UNIV DENMARK, DEPT MICROBIOL, BLDG 301, DK-2800 LYNGBY, DENMARK (Reprint); TECH UNIV DENMARK, DEPT MICROBIOL, DK-2800 LYNGBY, DENMARK; AARHUS UNIV, DEPT MED MICROBIOL, DK-8000 AARHUS C, DENMARK CYA DENMARK MICROBIOLOGY-UK, (JUN 1997) Vol. 143, Part 6, pp. 2027-2038. Publisher: SOC GENERAL MICROBIOLOGY, HARVEST HOUSE 62 LONDON ROAD, READING, BERKS, ENGLAND RG1 5AS. ISSN: 1350-0872. DTArticle; Journal FS LIFE LA English REC Reference Count: 34 *ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS* L7 ANSWER 15 OF 21 USPATFULL AB In order to obtain a novel binding protein against a chosen target, DNA molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein.

AN 96:101466 USPATFULL

TI Directed evolution of novel binding proteins
IN Ladner, Robert C., Ijamsville, MD, United States
Guterman, Sonia K., Belmont, MA, United States
Roberts, Bruce L., Milford, MA, United States
Markland, William, Milford, MA, United States
Ley, Arthur C., Newton, MA, United States
Kent, Rachel B., Boxborough, MA, United States

PA Protein Engineering Corporation, Cambridge, MA, United States (U.S.

corporation)

PI US 5571698 19961105 AI US 1993-57667 19930618 (8)

DCD 20100629

RLI Continuation of Ser. No. US 1991-664989, filed on 1 Mar 1991, now patented, Pat. No. US 5223409 which is a continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned which is a continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, now abandoned

DT Utility FS Granted

EXNAM Primary Examiner: Ulm, John

LREP Cooper, Iver P.
CLMN Number of Claims: 83
ECL Exemplary Claim: 1

DRWN 16 Drawing Figure(s); 16 Drawing Page(s)

LN.CNT 15323

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 16 OF 21 USPATFULL

AΒ In order to obtain a novel binding protein against a chosen target, DNA molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein.

AN 95:29292 USPATFULL

TI Viruses expressing chimeric binding proteins

IN Ladner, Robert C., Ijamsville, MD, United States Guterman, Sonia K., Belmont, MA, United States Roberts, Bruce L., Milford, MA, United States Markland, William, Milford, MA, United States

Ley, Arthur C., Newton, MA, United States Kent, Rachel B., Boxborough, MA, United States Protein Engineering Corporation, Cambridge, MA, United States (U.S. PΑ corporation) PΙ US 5403484 19950404 US 1993-9319 19930126 (8) ΑI Division of Ser. No. US 1991-664989, filed on 1 Mar 1991, now patented, RIT Pat. No. US 5223409 which is a continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned which is a continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, now abandoned WO 1989-3731 PRAI 19890901 DTUtility FS Granted Primary Examiner: Hill, Jr., Robert J.; Assistant Examiner: Ulm, John D. EXNAM Cooper, Iver P. LREP Number of Claims: 49 CLMN ECL Exemplary Claim: 1 16 Drawing Figure(s); 16 Drawing Page(s) LN.CNT 14368 CAS INDEXING IS AVAILABLE FOR THIS PATENT. **L7** ANSWER 17 OF 21 USPATFULL AΒ In order to obtain a novel binding protein against a chosen target, DNA molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein. AN 93:52487 USPATFULL ΤI Directed evolution of novel binding proteins TN Ladner, Robert C., Ijamsville, MD, United States Guterman, Sonia K., Belmont, MA, United States Roberts, Bruce L., Milford, MA, United States Markland, William, Milford, MA, United States Ley, Arthur C., Newton, MA, United States Kent, Rachel B., Boxborough, MA, United States PΑ Protein Engineering Corp., Cambridge, MA, United States (U.S. corporation) PΙ US 5223409 19930629 US 1991-664989 AΤ 19910301 (7) RLT Continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned And a continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, now abandoned DT Utility FS Granted EXNAM Primary Examiner: Hill, Jr., Robert J.; Assistant Examiner: Ulm, John D. Cooper, Iver P. LREP Number of Claims: 66 CLMN ECL Exemplary Claim: 1 16 Drawing Figure(s); 16 Drawing Page(s) LN.CNT 15410 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 18 OF 21 USPATFULL L7 AΒ The present invention relates to recombinant vector/host systems which can direct the expression of foreign genes under the control of the Heliothis polyhedrin promoter. Using the systems of the present invention, a heterologous gene of interest can be expressed as an unfused peptide or protein, a fusion protein, or as a recombinant occlusion body which comprises crystallized polyhedrin fusion proteins bearing the heterologous gene product on the surface of or within the occlusion body. The recombinant proteins or occlusion bodies of the present invention have uses in vaccine formulations and immunoassays, as biological insecticides, and as expression systems for the production of foreign peptides or proteins. ΑN 91:66733 USPATFULL TI Heliothis expression systems IN Fraser, Malcolm J.; South Bend, IN, United States Rosen, Elliot D., South Bend, IN, United States Ploplis, Victoria A., South Bend, IN, United States American Biogenetic Science, Inc., Copiague, NY, United States (U.S. PΑ corporation) PΙ US 5041379 19910820 US 1988-168109 ΑI 19880314 (7) Continuation-in-part of Ser. No. US 1987-26499, filed on 16 Mar 1987, RLI DT Utility FS Granted Primary Examiner: Schwartz, Richard A.; Assistant Examiner: Peet, EXNAM Richard C. LREP Pennie & Edmonds CLMN Number of Claims: 15 ECL Exemplary Claim: 1 26 Drawing Figure(s); 25 Drawing Page(s) CAS INDEXING IS AVAILABLE FOR THIS PATENT. L7 ANSWER 19 OF 21 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE AB The K88 fimbriae of enterotoxigenic Escherichia coli are strongly immunogenic antigens that can be used to evoke protective immunity. To find out whether these fimbriae can be used as carriers for foreign epitopes, a high variable region present in the primary structure of the different K88 variants was replaced with five different heterologous epitopes to investigate to what extent these insertions affected the expression, assembly (biogenesis), stability and immunogenic properties of the resulting hybrid fimbriae. Amino acid residues 163-173, were replaced using site-directed in vitro mutagenesis and the hybrid fimbriae were tested for these aspects using ELISA, immunoelectronmicroscopy and immunoblotting. Replacement of this highly variable region did not affect the biosynthesis of fimbriae, although all mutations tested resulted in a reduced expression depending on the epitope inserted. Testing of the different hybrid fimbriae with a panel of monoclonal antibodies raised against the various K88 serotypes K88ab, K88ac and K88ad indicated that replacement of amino acid sequence 163-173 did not affect conserved or K88ab specific epitopes but the K88ac and K88ad specific conformation was lost. Immunization with hybrid fimbriae raises antibodies specific for the inserted heterologous epitopes. AN 1990:426380 BIOSIS DNBA90:87181 K88 FIMBRIAE AS CARRIERS OF HETEROLOGOUS ANTIGENIC DETERMINANTS. ΤI ΑU BAKKER D; VAN ZIJDERVELD F G; VAN DER VEEN S; OUDEGA B; DE GRAAF F K BIOLOGISCH LABORATORIUM, VRIJE UNIVERSITEIT, DE BOELELAAN 1087, 1081 HV CS

AMSTERDAM, NETHERLANDS.

SO MICROB PATHOG, (1990) 8 (5), 343-352. CODEN: MIPAEV. ISSN: 0882-4010. FS BA; OLD English LA ANSWER 20 OF 21 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE L7 Hypervariable regions (HRs) of the major subunit of F11 fimbriae AB were exploited for insertion of foreign epitopes. Two insertion vectors were created that contain a unique cloning site in HR1 or HR4 respectively. Several oligonucleotides, coding for antigenic determinants derived from different pathogens, were cloned in both insertion vectors. Hybrid fimbrial subunits were generally shown to be assembled in fimbriae when the length of the inserted peptide did not exceed 14 amino acids. The inserted peptides appeared to be exposed in the fimbrial content. One hybrid fimbrial protein induced detectable levels of antibodies against the inserted epitope if injected into mice. ΑN 1990:494282 BIOSIS DN BA90:122628 EXPRESSION OF FOREIGN EPITOPES IN P-FIMBRIAE TI OF ESCHERICHIA-COLI. VAN DIE I; VAN OOSTERHOUT J; VAN MEGEN I; BERGMANS H; HOEKSTRA W; ΑU ENGER-VALK B; BARTELING S; MOOI F DEP. MEDICAL CHEMISTRY, VRIJE UNIVERSITEIT, VAN DER BOECHORSTSTRAAT 7, CS 1007 MC AMSTERDAM, NETH. MOL GEN GENET, (1990) 222 (2-3), 297-303. SO CODEN: MGGEAE. ISSN: 0026-8925. FS BA; OLD LA English L7 ANSWER 21 OF 21 USPATFULL The present invention is directed to recombinant baculoviruses which AB encode fusion polyhedrin proteins capable of forming occlusion bodies containing foreign peptides. The recombinant baculoviruses of the invention are formed by insertion into or replacement of regions of the polyhedrin gene that are not essential for occlusion body formation, with foreign DNA fragments by recombinant DNA techniques. The recombinant occlusion bodies produced in accordance with the present invention have uses in vaccine formulations, immunoassays, immobilized enzyme reactions, as biological insecticides, and as expression vectors. AN89:80739 USPATFULL Recombinant baculovirus occlusion bodies in vaccines and ΤI biological insecticides IN Fraser, Malcolm J., South Bend, IN, United States Rosen, Elliot D., South Bend, IN, United States Ploplis, Victoria A., South Bend, IN, United States PA American Biogenetic Sciences, Inc., Copiague, NY, United States (U.S. corporation) PΙ US 4870023 19890926 ΑI US 1988-153736 19880208 (7)

RLI Continuation-in-part of Ser. No. US 1987-26498, filed on 16 Mar 1987, now abandoned which is a continuation-in-part of Ser. No. US 1987-26499, filed on 16 Mar 1987

DT Utility

FS Granted

EXNAM Primary Examiner: Wiseman, Thomas G.; Assistant Examiner: Seidman, Stephanie

LREP Pennie & Edmonds
CLMN Number of Claims: 51
ECL Exemplary Claim: 1

DRWN 28 Drawing Figure(s); 26 Drawing Page(s)

LN.CNT 3868

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

(FILE 'HOME' ENTERED AT 11:23:50 ON 30 OCT 2002)

FILE 'BIOSIS, CABA, CAPLUS, EMBASE, LIFESCI, MEDLINE, SCISEARCH, USPATFULL, JAPIO' ENTERED AT 11:24:04 ON 30 OCT 2002

L1	82 S	VAN DIE, IRMA/AU
L2	0 S	FOREGIN EPITOPE
L3	890 S	FOREIGN EPITOPE
T 4	1 0	T 1 ANTO T 2

L4 1 S L1 AND L3

L5 2 S L1 AND VACCINE

L6 0 S VAN OOSTERNOUT, JOOST/AU

25 S BERGMANS, HANS/AU

L8 1 S L7 AND L3

L7

L9 0 S L1 AND PROTECTIVE

L10 732 S L1 AND IMMUNIZ OR VACINA?

L11 14406 S L3 AND FIMBRIAE OR PILI

L12 1267 S L11 AND VACCINE

L13 17 S L12 AND PAPA

L14 13 DUP REM L13 (4 DUPLICATES REMOVED)

FILE 'STNGUIDE' ENTERED AT 11:36:57 ON 30 OCT 2002

FILE 'BIOSIS, CABA, CAPLUS, EMBASE, LIFESCI, MEDLINE, SCISEARCH, USPATFULL, JAPIO' ENTERED AT 11:37:49 ON 30 OCT 2002

L15 1 S L14 AND L3

ANSWER 1 OF 13 USPATFULL L14 The present invention provides bacterial immunogenic agents for AB administration to humans and non-human animals to stimulate an immune response. Also provided are methods for vaccination of mammalian species, especially human patients, with variants of the E. coli FimH protein, said variants being derived from different strains of E. coli, and to production of antibodies that protect the vaccine recipient against infection by pathogenic bacterial species. In another aspect the invention provides antibodies against such proteins and protein complexes that may be used as diagnostics and/or as protective/treatment agents for pathogenic bacterial species. A plasmid-based method of producing polypeptides, especially fused polypeptides, such as the complex of a bacterial chaperone and a bacterial adhesin, is also disclosed. 2002:272472 USPATFULL AN TI FimH adhesin proteins and methods of use Langermann, Solomon, Baltimore, MD, UNITED STATES IN Revel, Andrew, Dallas, TX, UNITED STATES Auguste, Christine, Germantown, MD, UNITED STATES Burlein, Jeanne, Springfield, VA, UNITED STATES PΙ US 2002150587 A1 20021017 ΑI US 2001-900575 A1 20010706 (9) PRAI US 2000-216750P 20000707 (60) DT Utility FS APPLICATION Alan J. Gran, Esq., c/o CARELLA, BYRNE, BAIN, GILFILLAN, CECCHI, STEWART LREP & OLSTEIN, 6 Becker Farm Road, Roseland, NJ, 07068 CLMN Number of Claims: 38 ECL Exemplary Claim: 1 DRWN 17 Drawing Page(s) LN.CNT 2728 L14ANSWER 2 OF 13 USPATFULL AB The invention relates to compositions for the induction of anti-IgE antibodies in order to prevent or inhibit IgE-mediated disorders. The compositions contain carriers foreign to the immunized human or animal coupled to polypeptides containing fragments of the IgE molecule. The fragment of the IgE molecule includes the constant CH1 and/or the CH4 domain of the IgE molecule. The composition is administered to humans or animals in order to induce antibodies specific for endogenous IgE antibodies. These induced anti-IgE antibodies reduce or eliminate the pool of free IgE in the serum. Since many allergic diseases are mediated by IgE, IgE-mediated disorders are ameliorated in treated mammals. AN 2002:265550 USPATFULL ΤI Compositions for inducing self-specific anti-IgE antibodies and uses thereof Bachmann, Martin F., Winterhur, SWITZERLAND IN Renner, Wolfgang A., Zurich, SWITZERLAND Cytos Biotechnology AG (non-U.S. corporation) PA PΙ US 2002146422 Α1 20021010 20010727 (9) US 2001-916230 AΙ A1 US 2000-221841P 20000728 (60) PRAI DT Utility APPLICATION FS STERNE, KESSLER, GOLDSTEIN & FOX PLLC, 1100 NEW YORK AVENUE, N.W., SUITE LREP 600, WASHINGTON, DC, 20005-3934 CLMN Number of Claims: 46 ECL Exemplary Claim: 1 DRWN No Drawings LN.CNT 2138 CAS INDEXING IS AVAILABLE FOR THIS PATENT. L14 ANSWER 3 OF 13 USPATFULL AB A method of producing pili and vaccines containing

pili are described using bacteria that express at least one immunogenic peptide in a PapA region that does not normally contain such a peptide. 2002:258441 USPATFULL ΑN TI Immunogenic pili presenting foreign peptides, their production O'Hanley, Peter, Washington, DC, UNITED STATES ΙN Denich, Kenneth, Edmonton, CANADA Schmidt, M. Alexander, Muenster, GERMANY, FEDERAL REPUBLIC OF PΙ US 2002142008 A1 20021003 US 2001-833079 20010412 (9) ΑI Α1 US 2000-196491P 20000412 (60) PRAI DT Utility FS APPLICATION FOLEY AND LARDNER, SUITE 500, 3000 K STREET NW, WASHINGTON, DC, 20007 LREP CLMN Number of Claims: 7 ECL Exemplary Claim: 1 DRWN 5 Drawing Page(s) LN.CNT 967 CAS INDEXING IS AVAILABLE FOR THIS PATENT. L14 ANSWER 4 OF 13 USPATFULL AΒ A protein construct comprising a pilus protein portion, preferably a structurally stabilized pilus-protein, and an additional, or effector, portion other than a pilus protein or chaperone and wherein said effector portion serves to stabilize the pilus protein portion and to confer a therapeutic activity, such as vaccine activity or anti-microbial or anticancer activity, on the protein construct is disclosed. Such effector portion commonly comprises a donor strand complementary segment capable of structurally stabilizing a pilus protein subunit and attaching the auxiliary portion to said subunit to form the pilus protein analog of the invention. Methods of using said protein constructs are also disclosed as well as the formation and use of analogs comprising fragments of a pilus protein linked to effector components to produce immunogenic and/or therapeutic activity. AN 2002:164423 USPATFULL TТ Therapeutic compounds structurally-linked to bacterial polypeptides Hultgren, Scott J., Town and Country, MO, UNITED STATES IN Langermann, Solomon, Baltimore, MD, UNITED STATES Sauer, Frederic G., St. Louis, MO, UNITED STATES PΤ US 2002086037 A1 20020704 US 2001-27350 AΙ Α1 20011228 (10) US 2000-257880P PRAI 20001222 (60) DT Utility APPLICATION FS CARELLA, BYRNE, BAIN, GILFILLAN, CECCHI,, STEWART & OLSTEIN, 6 Becker LREP Farm Road, Roseland, NJ, 07068 Number of Claims: 65 CLMN ECL Exemplary Claim: 1 DRWN 10 Drawing Page(s) LN.CNT 1706 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 5 OF 13 USPATFULL L14AB The present invention relates to novel genes located in two chromosomal regions within uropathogenic E. coli that are associated with virulence. These chromosomal regions are known as pathogenicity islands (PAIs). In particular, the present application discloses 142 sequenced fragments (contigs) of DNA from two pools of cosmids covering pathogenicity islands PAI IV and PAI V located on the chromosome of the uropathogenic Escherichia coli J96. Further disclosed are 351 predicted protein-coding open reading frames within the sequenced fragments. 2002:141608 USPATFULL AN ΤI Nucleotide sequence of Escherichia coli pathogenicity islands

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Dillon, Patrick J., Carlsbad, CA, UNITED STATES
IN
       Choi, Gil H., Rockville, MD, UNITED STATES
       Welch, Rodney A., Madison, WI, UNITED STATES
       Human Genome Sciences, Inc., Rockville, MD, UNITED STATES (U.S.
PΑ
       corporation)
PΙ
       US 2002072595
                          A1
                               20020613
                               20010920 (9)
ΑI
       US 2001-956004
                          A1
       Division of Ser. No. US 1997-976259, filed on 21 Nov 1997, GRANTED, Pat.
RLI
       No. US 6316609
       US 1997-61953P
                           19971014 (60)
PRAI
                           19961122 (60)
       US 1996-31626P
DT
       Utility
FS
       APPLICATION
LREP
       HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, ROCKVILLE, MD, 20850
CLMN
       Number of Claims: 33
ECL
       Exemplary Claim: 1
DRWN
       2 Drawing Page(s)
LN.CNT 8481
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L14
    ANSWER 6 OF 13 USPATFULL
       A method of producing pili and vaccines containing
AB
       pili is described using bacteria harboring mutations that
      facilitate detachment of pili from the bacteria. Wild type
       pili have known immunoprotective effects in treating urinary
       tract infections. The mutant pili produced by this method are
       also shown to have such immunoprotective effects. Therefore, the
       pili may be used to make vaccines for treating urinary
       tract infections.
AN
       2002:105686 USPATFULL
ΤI
       Dissociated pili, their production and use
TN
       O'Hanley, Peter, Washington, DC, UNITED STATES
       Denich, Kenneth, Edmonton, CANADA
       US 2002054888
PΙ
                               20020509
                          Α1
       US 2001-833067
                               20010412 (9)
AΙ
                          A1
       US 2000-196493P
                           20000412 (60)
PRAI
DT
       Utility
       APPLICATION
FS
       Stephen B. Maebius, FOLEY & LARDNER, Suite 500, 3000 K Street, N.W.,
LREP
       Washington, DC, 20007-5109
CLMN
       Number of Claims: 5
ECL
       Exemplary Claim: 1
DRWN
       8 Drawing Page(s)
LN.CNT 727
L14 ANSWER 7 OF 13 CAPLUS COPYRIGHT 2002 ACS
     A the authors disclose the prepn. and isolation of pili from
     Escherichia coli with deletional mutations in papH. In a mouse model of
     pyelonephritis, vaccination with these pili prevented renal
     colonization. In addn., the authors disclose epitopes of papA
     and the use of these immunogenic peptide in a PapA region that
     does not normally contain such a peptide.
ΑN
     2001:780956 CAPLUS
DN
     135:343274
TI
     Immunogenic pili presenting foreign peptides: vaccination
     against urinary tract infections
IN
     Denich, Kenneth; Schmidt, M. Alexander
PA
     O'Hanley, Peter, USA
SO
     PCT Int. Appl., 35 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO. DATE
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                       A2
                            20011025
                                           WO 2001-US11918 20010412
ΡI
    WO 2001079277
     WO 2001079277
                      A3
                            20020523
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
       RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     US 2002142008
                       A1
                            20021003
                                          US 2001-833079
                                                           20010412
PRAI US 2000-196491P
                       Р
                            20000412
L14 ANSWER 8 OF 13 USPATFULL
       The present invention relates to novel genes located in two chromosomal
AB
       regions within uropathogenic E. coli that are associated with virulence.
       These chromosomal regions are known as pathogenicity islands (PAIs). In
       particular, the present application discloses 142 sequenced fragments
       (contigs) of DNA from two pools of cosmids covering pathogenicity
       islands PAI IV and PAI V located on the chromosome of the uropathogenic
       Escherichia coli J96. Further disclosed are 351 predicted protein-coding
       open reading frames within the sequenced fragments.
ΑN
       2001:202784 USPATFULL
TΙ
       Nucleotide sequence of Escherichia coli pathogenicity islands
       Dillon, Patrick J., Gaithersburg, MD, United States
ΤN
       Choi, Gil H., Rockville, MD, United States
       Welch, Rodney A., Madison, WI, United States
       Human Genome Sciences, Inc., Rockville, MD, United States (U.S.
PΑ
       corporation)
       Wisconsin Alumni Research Foundation, Madison, WI, United States (U.S.
       corporation)
PΙ
       US 6316609
                          B1
                               20011113
       US 1997-976259
AΙ
                               19971121 (8)
                           19971014 (60)
PRAI
       US 1997-61953P
       US 1996-31626P
                           19961122 (60)
DT
       Utility
FS
       GRANTED
      Primary Examiner: Clark, Deborah J. R.; Assistant Examiner: Sorbello,
EXNAM
       Eleanor
LREP
       Human Genome Sciences, Inc.
CLMN
       Number of Claims: 113
ECL
       Exemplary Claim: 1
DRWN
       2 Drawing Figure(s); 2 Drawing Page(s)
LN.CNT 3533
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
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L14 ANSWER 9 OF 13 USPATFULL

AΒ The DegP (HtrA) protease is a multifunctional protein essential for the removal of misfolded and aggregated proteins in the periplasm. The present invention provides an assay for inhibitors of DegP activity, comprising mixing a suspected inhibitor of DegP activity with DegP and a suitable substrate (preferably a native substrate of DegP such as PapA) and detecting changes in DegP activity. DegP has been shown to be essential for virulence in several Gram negative pathogens. Only three natural targets for DegP have been described: colicin A lysis protein (Cal), pilin subunits (K88, K99, Pap) and recently HMW1 and HMW2 from Hemophilus influenzae. In vitro, DegP has shown weak protease activity on casein and several other non-native substrates. The present inventors have identified the major pilin subunit of the Pap pilus, PapA, as a native DegP substrate and demonstrated binding and proteolysis of this substrate in vitro. Using an NH.sub.2 -terminal affinity tag the present inventors have purified PapA away

from the PapD chaperone, in the presence of denaturant, to use as a proteolysis substrate. This finding will allow the identification of the DegP recognition and cleavage sites in substrate proteins, and further, allow the design of small molecule inhibitors of protease function. AN 2001:185058 USPATFULL ΤI DeqP periplasmic protease a new anti-infective target and an in vitro assay for DegP protease function Jones, Hal C., Corvallis, OR, United States TN Liu, Christopher, Cambridge, MA, United States Hultgren, Scott J., Town and Country, MO, United States Hruby, Dennis E., Albany, OR, United States Franke, Christine A., Albany, OR, United States Evans, Amy K., West Linn, OR, United States Washington University, St. Louis, MO, United States (U.S. corporation) PΑ Siga Pharmaceuticals, New York, NY, United States (U.S. corporation) PΙ US 6306619 В1 20011023 US 2000-605858 20000629 (9) AΤ DTUtility FS GRANTED Primary Examiner: Mosher, Mary E. EXNAM Burns, Doane, Swecker & Mathis, L.L.P. LREP Number of Claims: 8 CLMN Exemplary Claim: 1 ECL 12 Drawing Figure(s); 8 Drawing Page(s) DRWN LN.CNT 615 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 10 OF 13 USPATFULL L14 An antigen which, as its major immunizing component, comprises a AB determinant of an adhesin polypeptide or an immunogenically active subsequence thereof or a precursor therefor which is convertible to an immunogenically active form, antibodies against which determinant react with the adhesin polypeptide produced by pathogenic adhesin-forming bacteria which adhere to mammalian tissue, antibodies against such antigen, and DNA expressing, as a principal gene product thereof, such antigen. 2001:158467 USPATFULL AN TI Anti-bodies binding adhesin-derived antigens Lindberg, Frederik Carl, Sandviken, Sweden IN Lund, Bjorn Olof, Umea, Sweden Baga, Britt Monika, Umea, Sweden Norgen, Mari Elisabet, Umea, Sweden Goransson, Mikael, Umea, Sweden Uhlin, Bernt Eric, Umea, Sweden Normark, Jan Staffan, Holmsund, Sweden Lark, David Lee, Umea, Sweden PA Symbicom Aktiebolag, Umea, Sweden (non-U.S. corporation) PT US 6291649 В1 20010918 US 1998-75396 19980511 (9) ΑI RLI Division of Ser. No. US 1995-447685, filed on 23 May 1995, now patented, Pat. No. US 5804198 Continuation of Ser. No. US 1993-123032, filed on 20 Sep 1993, now abandoned Continuation of Ser. No. US 1992-856829, filed on 23 Mar 1992, now abandoned Continuation of Ser. No. US 1991-678167, filed on 28 Mar 1991, now abandoned Continuation of Ser. No. US 1988-245469, filed on 16 Sep 1988, now abandoned Continuation of Ser. No. US 817849 DK 1984-2190 PRAI 19840502 DT Utility FS GRANTED EXNAM Primary Examiner: Graser, Jennifer E. Cooper, Iver P. LREP Number of Claims: 45 CLMN ECL Exemplary Claim: 1 DRWN 3 Drawing Figure(s); 3 Drawing Page(s)

LN.CNT 2145 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 11 OF 13 USPATFULL L14 An antigen which, as its major immunizing component, comprises a AB determinant of an adhesin polypeptide or an immunogenically active subsequence thereof or a precursor therefor which is convertible to an immunogenically active form, antibodies against which determinant react with the adhesin polypeptide produced by pathogenic adhesin-forming bacteria which adhere to mammalian tissue, antibodies against such antigen, and DNA expressing, as a principal gene product thereof, such antigen. 1998:108037 USPATFULL ANΤI Vaccines against disease caused by pathogenic pilus-forming bacteria Lindberg, Frederik Carl, Sandviken, Sweden IN Lund, Bjorn Olof, Ume.ang., Sweden B.ang.ga, Britt Monika, Ume.ang., Sweden Norgren, Mari Elisabet, Ume.ang., Sweden Goransson, Mikael, Ume.ang., Sweden Uhlin, Bernt Eric, Ume.ang., Sweden Normark, Jan Staffan, Holmsund, Sweden Lark, David Lee, Ume.ang., Sweden Symbicom Aktiebolag, Umea, Sweden (non-U.S. corporation) PΑ PΙ US 5804198 19980908 19950523 (8) ΑI US 1995-447685 Continuation of Ser. No. US 1993-123032, filed on 20 Sep 1993, now RLI abandoned which is a continuation of Ser. No. US 1992-856829, filed on 23 Mar 1992, now abandoned which is a continuation of Ser. No. US 1991-678167, filed on 28 Mar 1991, now abandoned which is a continuation of Ser. No. US 1988-245469, filed on 16 Sep 1988, now abandoned which is a division of Ser. No. US 1986-817849, filed on 19 Feb 1986, now patented, Pat. No. US 4795803 DK 1984-2190 PRAI 19840502 Utility DT FS Granted EXNAM Primary Examiner: Sidberry, Hazel F. Cooer, Iver P. LREP Number of Claims: 38 CLMN Exemplary Claim: 1 ECL 3 Drawing Figure(s); 3 Drawing Page(s) DRWN LN.CNT 2188 CAS INDEXING IS AVAILABLE FOR THIS PATENT. L14

ANSWER 12 OF 13 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE

Pyelonephritis-associated pili (Pap) are important in the AΒ pathogenesis of ascending, unobstructive Escherichia coli-caused renal infections because these surface bacterial organelles mediate diagalactoside-specific binding to host uroepithelial cells. Pap are composed of many different polypeptides, of which only the tip proteins mediate specific binding. The PapA moiety polymerizes to form the bulk of the pilus structure and has been employed in vaccines despite its lack of Gal.alpha.(1-4)Gal receptor specificity. Animal recipients of PapA pilus-based vaccines are protected against experimental pyelonephritis caused by homologous and heterologous Gal-Gal-binding uropathogenic E. coli strains. Specific PapA immunoglobulin G antibodies in urine are correlated with protection in these infection models. The nucleotide sequences of the gene encoding PapA were determined for three E. coli clones expressing F71, F72, and F9 pili and were compared with corresponding sequences for other F serotypes. Specific rabbit antisera were employed in enzyme-linked immunosorbent assays to study the cross-reactivity between Gal-Gal pili purified from recombinant strains expressing F71, F72, F9, or

F13 pili and among 60 Gal-Gal-binding wild-type strains. We present data which corroborate the concept that papA genes are highly homologous and encode proteins which exhibit > 70% homology among pili different serotypes. The differences primarily occur in the cysteine-cysteine loop and variable regions and constitute the basis for serological diversity of these pili. Although there are differences in primary structures among these pili, antisera raised against pili of one serotype cross-reacted frequently with many other Gal-Gal pili of different serotypes. Furthermore, antisera raised against pili of the F13 serotype cross-reacted strongly or moderately with 52 (86%) of 60 wild-type Gal-Gal-binding E. coli strains. These data suggest that there are common immunogenic domains among these proteins. These additional data further support the hypothesis that broadly cross-protective PapA pilus vaccines for the immunoprophylaxis of pyelonephritis might be developed. 1992:28956 BIOSIS BA93:18231 DNA SEQUENCES OF THREE PAPA GENES FROM UROPATHOGENIC ESCHERICHIA-COLI STRAINS EVIDENCE OF STRUCTURAL AND SEROLOGICAL CONSERVATION. DENICH K; BLYN L B; CRAIU A; BRAATEN B A; HARDY J; LOW D A; O'HANLEY P D DEP. MICROBIOLOGY IMMUNOLOGY, STANFORD UNIVERSITY, STANFORD, CALIF. 94305. INFECT IMMUN, (1991) 59 (11), 3849-3858. CODEN: INFIBR. ISSN: 0019-9567. BA; OLD English ANSWER 13 OF 13 USPATFULL An antigen which, as its major immunizing component, comprises a determinant of an adhesin polypeptide or an immunogenically active subsequence thereof or a precursor therefor which is convertible to an immunogenically active form, antibodies against which determinant react with the adhesion polypeptide produced by pathogenic adhesin-forming bacteria which adhere to mammalian tissue, antibodies against such antigen, and DNA expressing, as a principal gene product thereof, such antigen. 89:1283 USPATFULL Adhesin antigens, antibodies and DNA fragment encoding the antigen, methods and means for diagnosis and immunization etc. Lindberg, Frederick C., Sandviken, Sweden Lund, Bjorn O., Umea, Sweden Baga, Britt M., Umea, Sweden Norgren, Mari E., Umea, Sweden Goransson, Mikael, Umea, Sweden Uhlin, Bernt E., Umea, Sweden Normark, Jan S., Holmsund, Sweden Lark, David L., Umea, Sweden Syn-Tek AB, Umea, Sweden (non-U.S. corporation) US 4795803 19890103 WO 8505037 19851121 US 1986-817849 19860219 (6) WO 1985-DK45 19850502 19860219 PCT 371 date 19860219 PCT 102(e) date PRAI DK 1984-2190 19840502 Utility Granted Primary Examiner: Warden, Robert J.; Assistant Examiner: Saunders, David **EXNAM** LREP White, John P. Number of Claims: 10 CLMN ECL Exemplary Claim: 1 3 Drawing Figure(s); 3 Drawing Page(s) DRWN

ΑN DN

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LN.CNT 1912 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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ANSWER 1 OF 24 USPATFULL L6 In order to obtain a novel binding protein against a chosen target, DNA AB molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein. 2002:272761 USPATFULL AN ΤI Directed evolution of novel binding proteins TN Ladner, Robert Charles, Ijamsville, MD, UNITED STATES Guterman, Sonia Kosow, Belmont, MA, UNITED STATES Roberts, Bruce Lindsay, Milford, MA, UNITED STATES Markland, William, Milford, MA, UNITED STATES Ley, Arthur Charles, Newton, MA, UNITED STATES Kent, Rachel Baribault, Boxborough, MA, UNITED STATES PΙ US 2002150881 A1 20021017 ΑI US 2001-781988 20010214 (9) Α1 RLI Continuation of Ser. No. US 1998-192067, filed on 16 Nov 1998, ABANDONED Continuation of Ser. No. US 1995-415922, filed on 3 Apr 1995, PATENTED Continuation of Ser. No. US 1993-9319, filed on 26 Jan 1993, PATENTED Division of Ser. No. US 1991-664989, filed on 1 Mar 1991, PATENTED Continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, ABANDONED Continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, ABANDONED WO 1989-US3731 PRAI 19890901 Utility DT FS APPLICATION BROWDY AND NEIMARK, P.L.L.C., 624 Ninth Street, N.W., Washington, DC, LREP 20001 Number of Claims: 18 CLMN ECL Exemplary Claim: 1 DRWN 16 Drawing Page(s) LN.CNT 15696 CAS INDEXING IS AVAILABLE FOR THIS PATENT. L6 ANSWER 2 OF 24 USPATFULL A method of producing pili and vaccines containing AB pili are described using bacteria that express at least one immunogenic peptide in a PapA region that does not normally contain such a peptide. AN 2002:258441 USPATFULL Immunogenic pili presenting foreign peptides, their production ΤI IN O'Hanley, Peter, Washington, DC, UNITED STATES Denich, Kenneth, Edmonton, CANADA Schmidt, M. Alexander, Muenster, GERMANY, FEDERAL REPUBLIC OF PΤ US 2002142008 Α1 20021003 AΙ US 2001-833079 Α1 20010412 (9) PRAI US 2000-196491P 20000412 (60) DTUtility FS APPLICATION FOLEY AND LARDNER, SUITE 500, 3000 K STREET NW, WASHINGTON, DC, 20007 LREP CLMN Number of Claims: 7

Exemplary Claim: 1 ECL DRWN 5 Drawing Page(s) LN.CNT 967 CAS INDEXING IS AVAILABLE FOR THIS PATENT. L6 ANSWER 3 OF 24 USPATFULL The invention features method of inhibiting angiogenesis in a subject. AB The method includes decreasing syndecan-4 activity or expression in a cell, tissue, or subject. 2002:92068 USPATFULL AN Methods of modulating wound healing and angiogenesis ΤI Goetinck, Paul F., Boston, MA, UNITED STATES IN PΙ US 2002048585 **A1** 20020425 US 2001-900288 Α1 20010706 (9) AΙ 20000706 (60) PRAI US 2000-216247P DTUtility FS APPLICATION DIANA M. COLLAZO, Fish & Richardson P.C., 225 Franklin Street, Boston, LREP MA, 02110-2804 CLMN Number of Claims: 16 ECL Exemplary Claim: 1 DRWN 3 Drawing Page(s) LN.CNT 1486 CAS INDEXING IS AVAILABLE FOR THIS PATENT. 1.6 ANSWER 4 OF 24 USPATFULL AB The invention provides a highly efficient, rapid, and cost effective method of linking nucleic acid components in a predetermined order to produce a nucleic acid multicomponent construct. The invention further provides nucleic acid components, each nucleic acid component comprising a double stranded nucleic acid molecule having at least one single stranded 5' or 3' terminal sequence, the terminal sequence having sufficient complementarity to either a terminal sequence in a separate nucleic acid component or to a sequence in a linking nucleic acid molecule so as to allow for specific annealing of complementary sequences and linkage of the components in a predetermined order. Kits containing reagents required to practice the method of the invention are also provided. ΑN 2002:48253 USPATFULL METHOD AND KITS FOR PREPARING MULTICOMPONENT NUCLEIC ACID CONSTRUCTS TI HARNEY, PETER D., ALISO VIEJO, CA, UNITED STATES IN HARNEY, JENNIFER, ALISO VIEJO, CA, UNITED STATES PΤ US 2002028444 Α1 20020307 19981224 (9) US 1998-220398 AΙ Α1 Continuation-in-part of Ser. No. US 1997-877034, filed on 16 Jun 1997, RLI GRANTED, Pat. No. US 6277632 A 371 of International Ser. No. WO 1997-US10523, filed on 16 Jun 1997, UNKNOWN US 1996-19869P PRAI 19960617 (60) DTUtility FS APPLICATION ROPES & GRAY, ONE INTERNATIONAL PLACE, BOSTON, MA, 02110-2624 LREP Number of Claims: 42 CLMN ECL Exemplary Claim: 1 DRWN 5 Drawing Page(s) LN.CNT 3518 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 5 OF 24 USPATFULL 1.6 AB A strategically modified hepatitis B core protein is described, where an insert is provided, preferably in an immunodominant region of the nucleocapsid protein, containing a chemically reactive amino acid residue. The modified hepatitis B core protein or its aggregated

nucleocapsid protein particles can be pendently linked to a hapten to form a modified nucleocapsid conjugate. Such a conjugate is useful in

the preparation of vaccines or antibodies. The modified hepatitis B core protein can also be modified to include a T cell epitope. 2001:71101 USPATFULL AN Strategically modified hepatitis B core proteins and their derivatives TI Birkett, Ashley J., Solana Beach, CA, United States IN Immune Complex Corporation, San Diego, CA, United States (U.S. PA corporation) US 6231864 B1 20010515 PI US 1999-248588 19990211 (9) AΤ US 1998-74537P 19980212 (60) PRAI Utility DΤ FS Granted Primary Examiner: Wortman, Donna C. EXNAM Welsh & Katz, Ltd. LREP CLMN Number of Claims: 22 Exemplary Claim: 1 ECL1 Drawing Figure(s); 1 Drawing Page(s) DRWN LN.CNT 1665 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 6 OF 24 USPATFULL L6 Transgenically produced prolactin and methods of making and using AB transgenically produced prolactin are disclosed. 2001:47611 USPATFULL AN Transgenically produced prolactin TI Echelard, Yann, Brookline, MA, United States IN Wilburn, Brian, Boston, MA, United States Genzyme Transgenics Corporation, Framingham, MA, United States (U.S. PA corporation) PΙ US 6210736 В1 20010403 US 1998-94781 ΑI 19980615 (9) PRAI US 1997-49856P 19970617 (60) DTUtility FS Granted Primary Examiner: Hauda, Karen M.; Assistant Examiner: Shukla, Ram R. EXNAM LREP Fish & Richardson P.C. CLMN Number of Claims: 6 ECL Exemplary Claim: 1 DRWN No Drawings LN.CNT 1936 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 7 OF 24 CAPLUS COPYRIGHT 2002 ACS AB A method of generating chimeric genes encoding a fusion product of the agfA fimbrin and a foreign protein, such as an antigen, in a Salmonella host by chromosomal gene replacement is described. One embodiment of the invention is exemplified by the expression of a model epitope (PT3) obtained from the GP63 protein of Leishmania major, by formation of recombinant agfA genes encoding PT3 fusing proteins recombined at 10 different sites throughout the agfA gene. These fusions are shown to be expressed in the thin aggregative fimbriae on the surface of bacterial cell. The AgfA fimbrin of Salmonella (CsgA for E. coli) provides a flexible and stable vehicle for the expression of foreign epitopes in enterobacteriaceae and the subsequent thin aggregative fimbrae (curli) expression product provide an ideal organelle for presentation of the foreign epitopes at the cell surface. 2000:725786 CAPLUS AN 133:306338 DN TI Use of the agfA fimbrin of Salmonella to present foreign proteins on the surface of a bacterial host White, Aaron P.; Doran, James L.; Collison, S. Karen; Kay, William W. IN

Innovation and Development Corporation, University of Victoria, Can.

PA

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SO
    PCT Int. Appl., 139 pp.
     CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
     PATENT NO.
                     KIND DATE
                                           APPLICATION NO. DATE
                     ____
                                           WO 2000-CA356
                                                            20000405
PΙ
    WO 2000060102
                      A2
                            20001012
     WO 2000060102
                      Α3
                            20010104
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
             CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
             ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
             SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA,
             ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
             DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
             CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                            19990405
PRAI US 1999-127888P
                      Ρ
L6
     ANSWER 8 OF 24 USPATFULL
       The present invention is directed to recombinant genes and their encoded
AB
       proteins which are recombinant flagellin fusion proteins. Such fusion
       proteins comprise amino acid sequences specifying an epitope encoded by
       a flagellin structural gene and an epitope of a heterologous organism
       which is immunogenic upon introduction of the fusion protein into a
       vertebrate host. The recombinant genes and proteins of the present
       invention can be used in vaccine formulations, to provide
       protection against infection by the heterologous organism, or to provide
       protection against conditions or disorders caused by an antigen of the
       organism. In a specific embodiment, attenuated invasive bacteria
       expressing the recombinant flagellin genes of the invention can be used
       in live vaccine formulations. The invention is illustrated by
       way of examples in which epitopes of malaria circumsporozoite antigens,
       the B subunit of Cholera toxin, surface and presurface antigens of
       Hepatitis B. VP7 polypeptide of rotavirus, envelope glycoprotein of HIV,
       and M protein of Streptococcus, are expressed in recombinant flagellin
       fusion proteins which assemble into functional flagella, and which
       provoke an immune response directed against the heterologous epitope, in
       a vertebrate host.
       2000:134749 USPATFULL
ΑN
       Recombinant flagellin vaccines
ΤI
       Majarian, William R., Mt. Royal, NJ, United States
ΙN
       Stocker, Bruce A. D., Palo Alto, CA, United States
       Newton, Salete M. C., Mountain View, CA, United States
       American Cyanamid Company, Madison, NJ, United States (U.S. corporation)
PΑ
       The Board of Trustees of the Leland Stanford Junior University,
       Stanford, CA, United States (U.S. corporation)
       US 6130082
PΤ
                               20001010
ΑI
       US 1992-837668
                               19920214 (7)
       Continuation of Ser. No. US 1989-348430, filed on 5 May 1989, now
RLI
       abandoned which is a continuation-in-part of Ser. No. US 1988-190570,
       filed on 5 May 1988, now abandoned
DT
       Utility
FS
       Granted
EXNAM
       Primary Examiner: Mosher, Mary E.
       Hamilton, Brook, Smith & Reynolds, P.C.
LREP
       Number of Claims: 3
CLMN
ECL
       Exemplary Claim: 1
DRWN
       15 Drawing Figure(s); 17 Drawing Page(s)
LN.CNT 2404
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
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A CS31A protein capsule subunit having an aminoacid sequence modified by
AB
       at least one heterologous peptide, the CS31A protein capsule comprising
       said subunit, and micro-organisms having the CS31A protein capsule with
       its subunit aminoacid sequence modified by at least one heterologous
       peptide, are disclosed. Methods for preparing said subunits, CS31A
       protein capsules comprising same, and micro-organisms having CS31A
       protein capsules, as well as the use thereof for preparing
       vaccines, producing peptides and preparing immunoassays, are
       also disclosed.
       2000:98007 USPATFULL
ΑN
TI
       ClpG subunit of CS31A protein capsule containing heterologous peptides
IN
       Girardeau, Jean-Pierre, Saint Genes Champanelle, France
       Martin, Christine, La Roche Blanche, France
       Mechin, Marie-Claire, Beaumont, France
       Der Vartanian, Maurice, Saint Genes Champanelle, France
       Bousquet, Fran.cedilla.ois, Ceyrat, France
       Institut National de la Recherche Agronomique-INRA, Paris, France
PΑ
       (non-U.S. corporation)
       US 6096321
                               20000801
PΤ
       WO 9414967
                  19940707
       US 1996-491954
ΑI
                               19960216 (8)
       WO 1993-FR1281
                               19931221
                                         PCT 371 date
                               19960216
                               19960216 PCT 102(e) date
PRAI
       FR 1992-15464
                           19921222
DT
       Utility
FS
       Granted
       Primary Examiner: Chin, Christopher L.; Assistant Examiner: Ryan, V.
EXNAM
       Schnader Harrison Segal & Lewis LLP
LREP
CLMN
       Number of Claims: 29
ECL
       Exemplary Claim: 1
       61 Drawing Figure(s); 53 Drawing Page(s)
DRWN
LN.CNT 3468
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L6
     ANSWER 10 OF 24 USPATFULL
       The present invention is concerned with vaccination of mammals against
AB
       GnRH. The vaccine comprises a GnRH peptide conjugate to E.
       coli fimbrial-filaments and elicits an immune response against GnRH.
       2000:12446 USPATFULL
AN
ΤI
       Carrier system against GnRH
       Van Der Zee, Anna, Woerden, Netherlands
IN
       Van Die, Irma Marianne, Gouda, Netherlands
       Hoekstra, Willem Pieter Martin, Zeist, Netherlands
       Gielen, Josephus Theodorus, St. Antohonis, Netherlands
       Akzo Nobel N.V., Arnhem, Netherlands (non-U.S. corporation)
PA
PΙ
       US 6019983
                                20000201
ΑI
       US 1995-521079
                                19950829 (8)
       Continuation of Ser. No. US 1993-78661, filed on 16 Jun 1993, now
RLI
       abandoned
       NL 1982-92201775
PRAI
                           19820619
DT
       Utility
FS
       Granted
       Primary Examiner: Sidberry, Hazel F.
EXNAM
LREP
       Gormley, Mary E., Blackstone, William M.
CLMN
       Number of Claims: 6
       Exemplary Claim: 1
ECL
       9 Drawing Figure(s); 9 Drawing Page(s)
LN.CNT 1366
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
L6
     ANSWER 11 OF 24 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE
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Recombinant live oral vaccines expressing pathogen-derived

AB

antiqens offer a unique set of attractive properties. Among these are the simplicity of administration, the capacity to induce mucosal and systemic immunity, and the advantage of permitting genetic manipulation for optimal antigen presentation. In this study, the benefit of having a heterologous antigen expressed on the surface of a live vector rather than intracellularly was evaluated. Accordingly, the immune response of mice immunized with a Salmonella enterica serovar Typhimurium vaccine strain expressing the Escherichia coli 987P fimbrial antigen on its surface (Fas+) was compared with the expression in the periplasmic compartments (Fas-). Orally immunized BALB/c mice showed that 987P fimbriated Salmonella serovar Typhimurium CS3263 (aroA asd) with pCS151 (fas+ asd+) elicited a significantly higher level of 987P-specific systemic immunoglobulin G (IgG) and mucosal IgA than serovar Typhimurium CS3263 with pCS152 (fasD mutant, asd+) expressing 987P periplasmic antigen. Further studies were aimed at determining whether the 987P fimbriae expressed by serovar Typhimurium chi4550 (cya crp asd) could be used as carriers of foreign epitopes. For this, the vaccine strain was genetically engineered to express chimeric fimbriae carrying the transmissible gastroenteritis virus (TGEV) C (379-388) and A (521-531) epitopes of the spike protein inserted into the 987P major fimbrial subunit FasA. BALB/c mice administered orally serovar Typhimurium chi4550 expressing the chimeric fimbriae from the tet promoter in pCS154 (fas+ asd+) produced systemic antibodies against both fimbria and the TGEV C epitope but not against the TGEV A epitope. To improve the immunogenicity of the chimeric fimbriae, the in vivo inducible nirB promoter was inserted into pCS154, upstream of the fas genes, to create pCS155. In comparison with the previously used vaccine, BALB/c mice immunized orally with serovar Typhimurium chi4550/pCS155 demonstrated significantly higher levels of serum IgG and mucosal IgA against 987P fimbria. Moreover, mucosal IqA against the TGEV C epitope was only detected with serovar Typhimurium chi4550/pCS155. The induced antibodies also recognized the epitopes in the context of the full-length TGEV spike protein. Hence, immune responses to heterologous chimeric fimbriae on Salmonella vaccine vectors can be optimized by using promoters known to be activated in vivo.

- AN 2000:291353 BIOSIS
- DN PREV200000291353
- TI Mucosal and systemic immune responses to chimeric **fimbriae** expressed by Salmonella enterica serovar Typhimurium **vaccine** strains.
- AU Chen, Huaiqing; Schifferli, Dieter M. (1)
- CS (1) University of Pennsylvania School of Veterinary Medicine, 3800 Spruce St., Philadelphia, PA, 19104-6049 USA
- SO Infection and Immunity, (June, 2000) Vol. 68, No. 6, pp. 3129-3139. print. ISSN: 0019-9567.
- DT Article
- LA English
- SL English
- L6 ANSWER 12 OF 24 CAPLUS COPYRIGHT 2002 ACS
- AB A review, with 50 refs., discussing fimbrial display of foreign epitopes, heterologous antigen display, antigen display by other fimbriae, and random library display.
- AN 2001:3230 CAPLUS
- DN 134:176932
- TI Fimbrial surface display systems in bacteria: from **vaccines** to random libraries
- AU Klemm, Per; Schembri, Mark A.
- CS Department of Microbiology, Technical University of Denmark, Lyngby, DK-2800, Den.
- SO Microbiology (Reading, United Kingdom) (2000), 146(12), 3025-3032 CODEN: MROBEO; ISSN: 1350-0872
- PB Society for General Microbiology

DT Journal; General Review

LA English

RE.CNT 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 13 OF 24 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 2

Objective: To construct the display vector based on the CS3 pili AB of enterotoxigenic Escherichia coli. Methods: The secondary structure antigen epitopes, hydrophilicity and flexibility of CS3 subunit were predicted with the Goldkey software. Based on the prediction, the site for inserting heterologous epitopes was chosen. Mutation was done using the overlapping extention PCR. The gene fragment coding for the VP1 of foot-mouth disease virus (FMDV) was synthesized and inserted into CS3. The surface expression of hybrid protein was examined using whole-cell ELISA, electron microscopy and immuno-electron microscopy. Mice were immunized by injecting the recombinant bacteria intraperitoneally to evaluate the immunogenicity of the hybrid proteins. Results: The VP1 of FMDV was displayed on the surface of the recombinant cells. The fusion proteins were expressed as hybrid pili. Mice produced antibody response against CS3 and the VP1 of FMDV. Conclusion: The CS3 pili can be a vector to express the foreign epitopes on the surface of the recombinant cells, and it may probably be an expression vector for the construction of the live gene engineering vaccine

AN 2001:49887 BIOSIS

DN PREV200100049887

TI Construction of a display vector based on the CS3 **pili** of enterotoxigenic Escherichia coli.

AU Gao Rongkai; Zhang Zhaoshan (1); Li Shuqin

CS (1) Academy of Military Medical Science, Institute of Biotechnology, Beijing, 100071: zhangzs@nic.bmi.ac.cn China

SO Zhonghua Weishengwuxue He Mianyixue Zazhi, (November, 2000) Vol. 20, No. 6, pp. 485-488. print. ISSN: 0254-5101.

DT Article

LA Chinese

SL Chinese; English

L6 ANSWER 14 OF 24 SCISEARCH COPYRIGHT 2002 ISI (R)

AΒ The display of peptide segments on the surface of bacteria offers many new and exciting applications in biotechnology and medical research. Fimbria-assisted display of heterologous sequences is a paradigm for chimeric organelle display on bacteria. Fimbriac are particularly attractive candidates for epitope display for several reasons: (1) they are present in extremely high numbers at the cell surface, (2) they are strong immunogens, (3) they possess inherent adhesive properties, and (4) they can be easily purified. The majority of work dealing with fimbria-assisted peptide display has been focused on the development of recombinant vaccines. A number of different fimbrial types have been used to display immune-relevant sectors of various foreign proteins. Chimeric fimbrial vaccines can be used in the context of purified proteins, however the potential also exists to exploit this technology for the development of live recombinant vaccines. Work has also bern performed demonstrating the amenability of fimbriae towards the powerful technology of random peptide display. This review summarises the current state of research in this field.

AN 2000:632511 SCISEARCH

GA The Genuine Article (R) Number: 344MZ

TI Fimbriae-assisted bacterial surface display of heterologous peptides

AU Klemm P (Reprint); Schembri M A

CS TECH UNIV DENMARK, DEPT MICROBIOL, BLDG 301, DK-2800 LYNGBY, DENMARK

(Reprint)
CYA DENMARK
SO INTERNATIONAL JOURNAL OF MEDICAL MICROBIOLOGY, (JUL 2000) Vol. 290, No. 3, pp. 215-221.
Publisher: URBAN & FISCHER VERLAG, BRANCH OFFICE JENA, P O BOX 100537, D-07705 JENA, GERMANY.

ISSN: 1438-4221.

DT Article; Journal

FS LIFE

LA English

REC Reference Count: 41

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

L6 ANSWER 15 OF 24 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 3

AB The strong immunogenicity of bacterial fimbriae results from their polymeric and proteinaceous nature, and the protective role of these immunogens in experimental or commercial vaccines is associated with their capacity to induce antiadhesive antibodies. Fimbria -mediated intestinal colonization by enteropathogens typically leads to similar antibody responses. The possibility of taking advantage of these properties was investigated by determining whether enteroadhesive fimbriae, like the 987P fimbriae of enterotoxigenic Escherichia coli, can serve as carriers for foreign antigens without losing their adhesive characteristics. Random linker insertion mutagenesis of the fasA gene encoding the major 987P subunit identified five different mutants expressing wild-type levels of fimbriation. The linker insertion sites of these mutants were used to introduce three continuous segments of viral surface glycoproteins known to be accessible to antibodies. These segments encode residues 11 to 19 or 272 to 279 of herpes simplex virus type 1 (HSV-1) glycoprotein D (gD(11-19) and gD(272-279), respectively) or residues 379 to 388 of the transmissible gastroenteritis virus (TGEV) spike protein (S(379-388)). Studies of bacteria expressing fimbriae incorporating mutated FasA subunits alone or together with wild-type FasA subunits (hybrid fimbriae) indicated that foreign epitopes were best exported and displayed on assembled fimbriae when they were inserted near the amino terminus of FasA. Fimbriated bacteria expressing FasA subunits carrying the HSV qD(11-19) or the TGEV S(379-388) epitope inserted between the second and third residues of mature FasA elicited high levels of foreign epitope antibodies in all rabbits immunized parenterally. Antibodies against the HSV epitope were also shown to recognize the epitope in the context of the whole gD protein. Because the 987P adhesive subunit FasG was shown to be present on mutated fimbriae and to mediate bacterial attachment to porcine intestinal receptors, polymeric display of foreign epitopes on 987P offers new opportunities to test the potential beneficial effect of enteroadhesion for mucosal immunization and protection against various enteric pathogens.

AN 1999:99340 BIOSIS

DN PREV199900099340

- TI Polymeric display of immunogenic epitopes from herpes simplex virus and transmissible gastroenteritis virus surface proteins on an enteroadherent fimbria.
- AU Rajini Rani, D. B.; Bayer, Manfred E.; Schifferli, Dieter M. (1)
- CS (1) Univ. Pa. Sch. Veterinary Med., 3800 Spruce St., Philadelphia, PA 19104-6049 USA
- SO Clinical and Diagnostic Laboratory Immunology, (Jan., 1999) Vol. 6, No. 1, pp. 30-40.
 ISSN: 1071-412X.
- DT Article
- LA English
- L6 ANSWER 16 OF 24 USPATFULL
- AB In order to obtain a novel binding protein against a chosen target, DNA

molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein. 1998:143904 USPATFULL Directed evolution of novel binding proteins Ladner, Robert Charles, Ijamsville, MD, United States Gutterman, Sonia Kosow, Belmont, MA, United States Roberts, Bruce Lindsay, Milford, MA, United States Markland, William, Milford, MA, United States Ley, Arthur Charles, Newton, MA, United States Kent, Rachel Baribault, Boxborough, MA, United States Dyax, Corp., Cambridge, MA, United States (U.S. corporation) US 5837500 19981117 US 1995-415922 19950403 (8) Continuation of Ser. No. US 1993-9319, filed on 26 Jan 1993, now patented, Pat. No. US 5403484 which is a division of Ser. No. US 1991-664989, filed on 1 Mar 1991, now patented, Pat. No. US 5223409 which is a continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned which is a continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, now abandoned Utility Granted EXNAM Primary Examiner: Ulm, John Cooper, Iver P. Number of Claims: 43 Exemplary Claim: 1 16 Drawing Figure(s); 16 Drawing Page(s) LN.CNT 15973 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 17 OF 24 USPATFULL The present invention is concerned with vaccination of mammals against GnRH. The vaccine comprises a GnRH peptide conjugate to E. coli fimbrial-filaments and elicits an immune response against GnRH. 97:101896 USPATFULL Carrier system against GNRH Van Der Zee, Anna, Woerden, Netherlands Van Die, Irma Marianne, Gouda, Netherlands Hoekstra, Willem Pieter Martin, Zeist, Netherlands Gielen, Josephus Theodorus, St. Antohonis, Netherlands AKZO Nobel N.V., Arnhem, Netherlands (non-U.S. corporation) US 5684145 19971104 US 1995-453588 19950530 (8) Division of Ser. No. US 1993-78661, filed on 16 Jun 1993, now abandoned NL 1992-1775 19920618 Utility Granted Primary Examiner: Sidberry, Hazel F. EXNAM Gormley, Mary E. Number of Claims: 8

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LREP

CLMN

ECL

L6

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RLI

PRAI

LREP

CLMN

Exemplary Claim: 1

ECL

RLI

DRWN 9 Drawing Figure(s); 9 Drawing Page(s) LN.CNT 1299 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 18 OF 24 USPATFULL

In order to obtain a novel binding protein against a chosen target, DNA AB molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein.

AN 96:101466 USPATFULL

Directed evolution of novel binding proteins
Ladner, Robert C., Ijamsville, MD, United States
Guterman, Sonia K., Belmont, MA, United States
Roberts, Bruce L., Milford, MA, United States
Markland, William, Milford, MA, United States
Ley, Arthur C., Newton, MA, United States
Kent, Rachel B., Boxborough, MA, United States

PA Protein Engineering Corporation, Cambridge, MA, United States (U.S.

corporation)

PI US 5571698 19961105 AI US 1993-57667 19930618 (8)

DCD 20100629

RLI Continuation of Ser. No. US 1991-664989, filed on 1 Mar 1991, now patented, Pat. No. US 5223409 which is a continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned which is a continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, now abandoned

DT Utility FS Granted

EXNAM Primary Examiner: Ulm, John

LREP Cooper, Iver P.
CLMN Number of Claims: 83
ECL Exemplary Claim: 1

DRWN 16 Drawing Figure(s); 16 Drawing Page(s)

LN.CNT 15323

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 19 OF 24 USPATFULL

AB In order to obtain a novel binding protein against a chosen target, DNA molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment,

the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein. 95:29292 USPATFULL AN Viruses expressing chimeric binding proteins TΤ Ladner, Robert C., Ijamsville, MD, United States TN Guterman, Sonia K., Belmont, MA, United States Roberts, Bruce L., Milford, MA, United States Markland, William, Milford, MA, United States Ley, Arthur C., Newton, MA, United States Kent, Rachel B., Boxborough, MA, United States Protein Engineering Corporation, Cambridge, MA, United States (U.S. PΑ corporation) PΙ US 5403484 19950404 US 1993-9319 19930126 (8) AΙ Division of Ser. No. US 1991-664989, filed on 1 Mar 1991, now patented, RLI Pat. No. US 5223409 which is a continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned which is a continuation-in-part of Ser. No. US 1988-240160, filed on 2 Sep 1988, now abandoned PRAI WO 1989-3731 19890901 DTUtility FS Granted EXNAM Primary Examiner: Hill, Jr., Robert J.; Assistant Examiner: Ulm, John D. Cooper, Iver P. LREP Number of Claims: 49 CLMN Exemplary Claim: 1 ECL 16 Drawing Figure(s); 16 Drawing Page(s) DRWN LN.CNT 14368 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 20 OF 24 USPATFULL

In order to obtain a novel binding protein against a chosen target, DNA AB molecules, each encoding a protein comprising one of a family of similar potential binding domains and a structural signal calling for the display of the protein on the outer surface of a chosen bacterial cell, bacterial spore or phage (genetic package) are introduced into a genetic package. The protein is expressed and the potential binding domain is displayed on the outer surface of the package. The cells or viruses bearing the binding domains which recognize the target molecule are isolated and amplified. The successful binding domains are then characterized. One or more of these successful binding domains is used as a model for the design of a new family of potential binding domains, and the process is repeated until a novel binding domain having a desired affinity for the target molecule is obtained. In one embodiment, the first family of potential binding domains is related to bovine pancreatic trypsin inhibitor, the genetic package is M13 phage, and the protein includes the outer surface transport signal of the M13 gene III protein.

AN 93:52487 USPATFULL

TI Directed evolution of novel binding proteins

IN Ladner, Robert C., Ijamsville, MD, United States Guterman, Sonia K., Belmont, MA, United States Roberts, Bruce L., Milford, MA, United States Markland, William, Milford, MA, United States Ley, Arthur C., Newton, MA, United States Kent, Rachel B., Boxborough, MA, United States

PA Protein Engineering Corp., Cambridge, MA, United States (U.S. corporation)

PI US 5223409 19930629 AI US 1991-664989 19910301 (7)

RLI Continuation-in-part of Ser. No. US 1990-487063, filed on 2 Mar 1990, now abandoned And a continuation-in-part of Ser. No. US 1988-240160,

filed on 2 Sep 1988, now abandoned DT Utility FS Granted EXNAM Primary Examiner: Hill, Jr., Robert J.; Assistant Examiner: Ulm, John D. Cooper, Iver P. LREP CLMN Number of Claims: 66 ECL Exemplary Claim: 1 16 Drawing Figure(s); 16 Drawing Page(s) DRWN CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 21 OF 24 USPATFULL L6 The present invention relates to recombinant vector/host systems which AΒ can direct the expression of foreign genes under the control of the Heliothis polyhedrin promoter. Using the systems of the present invention, a heterologous gene of interest can be expressed as an unfused peptide or protein, a fusion protein, or as a recombinant occlusion body which comprises crystallized polyhedrin fusion proteins bearing the heterologous gene product on the surface of or within the occlusion body. The recombinant proteins or occlusion bodies of the present invention have uses in vaccine formulations and immunoassays, as biological insecticides, and as expression systems for the production of foreign peptides or proteins. AN 91:66733 USPATFULL Heliothis expression systems TITN Fraser, Malcolm J., South Bend, IN, United States Rosen, Elliot D., South Bend, IN, United States Ploplis, Victoria A., South Bend, IN, United States American Biogenetic Science, Inc., Copiague, NY, United States (U.S. PA corporation) PΙ US 5041379 19910820 ΑI US 1988-168109 19880314 (7) Continuation-in-part of Ser. No. US 1987-26499, filed on 16 Mar 1987, RLI now abandoned DΤ Utility FS Granted EXNAM Primary Examiner: Schwartz, Richard A.; Assistant Examiner: Peet, Richard C. Pennie & Edmonds LREP Number of Claims: 15 CLMN Exemplary Claim: 1 ECL 26 Drawing Figure(s); 25 Drawing Page(s) DRWN LN.CNT 3494 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 22 OF 24 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. 1.6 The K88 fimbriae of enterotoxigenic Escherichia coli are AB strongly immunogenic antigens that can be used to evoke protective immunity. To find out whether these fimbriae can be used as carriers for foreign epitopes, a high variable region present in the primary structure of the different K88 variants was replaced with five different heterologous epitopes to investigate to what extent these insertions affected the expression, assembly (biogenesis), stability and immunogenic properties of the resulting hybrid fimbriae. Amino acid residues 163-173, were replaced using site-directed in vitro mutagenesis and the hybrid fimbriae were tested for these aspects using ELISA, immunoelectronmicroscopy and immunoblotting. Replacement of this highly variable region did not affect the biosynthesis of fimbriae, although all mutations tested resulted in a reduced expression depending on the epitope inserted. Testing of the different hybrid fimbriae with a panel of monoclonal antibodies raised against the various K88 serotypes K88ab, K88ac and K88ad indicated that replacement of amino acid sequence 163-173

did not affect conserved or K88ab specific epitopes but the K88ac and

K88ad specific conformation was lost. Immunization with hybrid **fimbriae** raises antibodies specific for the inserted heterologous epitopes.

AN 1990:426380 BIOSIS

DN BA90:87181

TI K88 FIMBRIAE AS CARRIERS OF HETEROLOGOUS ANTIGENIC DETERMINANTS.

AU BAKKER D; VAN ZIJDERVELD F G; VAN DER VEEN S; OUDEGA B; DE GRAAF F K

CS BIOLOGISCH LABORATORIUM, VRIJE UNIVERSITEIT, DE BOELELAAN 1087, 1081 HV AMSTERDAM, NETHERLANDS.

SO MICROB PATHOG, (1990) 8 (5), 343-352. CODEN: MIPAEV. ISSN: 0882-4010.

FS BA; OLD

LA English

L6 ANSWER 23 OF 24 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 4

AB Hypervariable regions (HRs) of the major subunit of F11 fimbriae were exploited for insertion of foreign epitopes. Two insertion vectors were created that contain a unique cloning site in HR1 or HR4 respectively. Several oligonucleotides, coding for antigenic determinants derived from different pathogens, were cloned in both insertion vectors. Hybrid fimbrial subunits were generally shown to be assembled in fimbriae when the length of the inserted peptide did not exceed 14 amino acids. The inserted peptides appeared to be exposed in the fimbrial content. One hybrid fimbrial protein induced detectable levels of antibodies against the inserted epitope if injected into mice.

AN 1990:494282 BIOSIS

DN BA90:122628

TI EXPRESSION OF FOREIGN EPITOPES IN P-FIMBRIAE OF ESCHERICHIA-COLI.

AU VAN DIE I; VAN OOSTERHOUT J; VAN MEGEN I; BERGMANS H; HOEKSTRA W; ENGER-VALK B; BARTELING S; MOOI F

CS DEP. MEDICAL CHEMISTRY, VRIJE UNIVERSITEIT, VAN DER BOECHORSTSTRAAT 7, 1007 MC AMSTERDAM, NETH.

SO MOL GEN GENET, (1990) 222 (2-3), 297-303. CODEN: MGGEAE. ISSN: 0026-8925.

FS BA; OLD

LA English

L6 ANSWER 24 OF 24 USPATFULL

The present invention is directed to recombinant baculoviruses which encode fusion polyhedrin proteins capable of forming occlusion bodies containing foreign peptides. The recombinant baculoviruses of the invention are formed by insertion into or replacement of regions of the polyhedrin gene that are not essential for occlusion body formation, with foreign DNA fragments by recombinant DNA techniques. The recombinant occlusion bodies produced in accordance with the present invention have uses in vaccine formulations, immunoassays, immobilized enzyme reactions, as biological insecticides, and as expression vectors.

AN 89:80739 USPATFULL

TI Recombinant baculovirus occlusion bodies in **vaccines** and biological insecticides

IN Fraser, Malcolm J., South Bend, IN, United States Rosen, Elliot D., South Bend, IN, United States Ploplis, Victoria A., South Bend, IN, United States

PA American Biogenetic Sciences, Inc., Copiague, NY, United States (U.S. corporation)

PI US 4870023 19890926

AI US 1988-153736 19880208 (7)

RLI Continuation-in-part of Ser. No. US 1987-26498, filed on 16 Mar 1987, now abandoned which is a continuation-in-part of Ser. No. US 1987-26499, filed on 16 Mar 1987

DT Utility FS Granted

EXNAM Primary Examiner: Wiseman, Thomas G.; Assistant Examiner: Seidman,

Stephanie

LREP Pennie & Edmonds
CLMN Number of Claims: 51
ECL Exemplary Claim: 1

DRWN 28 Drawing Figure(s); 26 Drawing Page(s)

LN.CNT 3868

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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(FILE 'HOME' ENTERED AT 12:01:09 ON 30 OCT 2002)

FILE 'AGRICOLA, LIFESCI, CONFSCI, BIOSIS, VETU, VETB, PHIN, PHIC' ENTERED AT 12:01:27 ON 30 OCT 2002

FILE 'BIOSIS, CABA, CAPLUS, EMBASE, LIFESCI, MEDLINE, SCISEARCH, USPATFULL, JAPIO' ENTERED AT 12:01:37 ON 30 OCT 2002

L1 644 S FOREIGN EPITOPES

L2 138412 S PILI OR FIMBRIAE OR FIMBRIA OR FIBRIN

L3 56 S L1 AND L2

L4 440609 S VACCINE

L5 36 S L3 AND L4

L6 24 DUP REM L5 (12 DUPLICATES REMOVED)

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